10/537767 JC17 Rec'd PCT/PTO 06 JUN 2005

96

SEQUENCE LISTING

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· <151> 2002-12-05

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<222> (1).. (1719)

<223> Inventor: Endo, Yuki; Endoh, Hideki; Ueda, Yoshitaka; Kato, Miyuki;

Inabe, Kazunori

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Met Ala Ala Val Pro Pro Leu Arg Asp Arg Leu Ser Phe Leu His Arg

1 10 15

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| _eu | Pro | | Leu 20 | Leu | Lys | Gly | Thr | Ser 25 | Asp _. | Asp | Ser | He | Pro 30 | Cys | Pro | |
|-------------------|-------------------|-------------------|-------------------|------------------|-------------------|-------------------|-------------------|-------------------|------------------|-------------------|-------------------|-------------------|-------------------|------------------|-------------------|-----|
| ggc Gly | Tyr | ctg Leu 35 | ttt Phe | gaa Glu | gag Glu | atc lle | gcc Ala 40 | aag Lys | att He | tcc Ser | cac His | gag Glu 45 | tca Ser | cta Leu | ggc Gly | 144 |
| agc Ser | agc Ser 50 | cag Gin | tgc Cys | ctg Leu | .ctg Leu | gag Glu 55 | tac Tyr | ctc Leu | ctg Leu | aac Asn | cgt Arg 60 | ctg Leu | gac Asp | agc Ser | agc Ser | 192 |
| tcc Ser 65 | ggc Gly | cac His | gtg Val | aag Lys | ctc Leu 70 | aag Lys | gtg Val | cta Leu | aag Lys | atc Ile 75 | ttg Leu | ctt Leu | tac Tyr | ctg Leu | tgt Cys 80 | 240 |
| ggt Gly | cat His | ggc Gly | tct Ser | tcc Ser 85 | tcc Ser | ttc Phe | ctc Leu | ctc Leu | atc He 90 | ctc Leu | agg Arg | aga Arg | aac Asn | tct Ser 95 | gct Ala | 288 |
| ctc Leu | atc Ile | caa GIn | gaa Glu 100 | gcc Ala | acg Thr | gct Ala | ttc Phe | tca Ser 105 | Gly | cct Pro | cca Pro | gat Asp | cct Pro 110 | Leu | cac His | 336 |
| gga Gly | aat Asn | agc Ser 115 | Leu | tac Tyr | cag Gln | aag Lys | gtg Val 120 | Arg | gcg | gct Ala | gcc Ala | cag Gln 125 | Asp | ctg Leu | ggt Gly | 384 |
| agc Ser | acc Thr 130 | ctg Leu | ttc Phe | tca Ser | gat Asp | gcc Ala 135 | Val | cca Pro | cag Gln | cct Pro | cca Pro 140 | Ser | cag Gln | cca Pro | cct Pro | 432 |
| cag GIn 145 | He | ccg Pro | cct Pro | ccc Pro | gca Ala 150 | Gly | atg Met | g ggo | gcc Ala | cag Gln 155 | n Ala | aga Arg | cct Pro | ctt Leu | agt Ser 160 | 480 |
| ~ 00 | 0± « | 000 | | ++ | | tac | : acs | , aa | gag | ago | : ago | Cgg | aca | ggc | tcc | 528 |

| Ala | Leu | Gln | Gly | Phe 165 | Gly | Tyr | Thr | Lys | G I บ 170 | Ser | Ser | Arg | Thr | Gly 175 | Ser | | |
|-----|-----|-----|-----|------------|-----|-----|-----|-----|--------------|-----|-----|-----|-----|-------------------|-----|---|-----|
| | | | | | | | | | | | | | | gta Val | | | 576 |
| _ | | | | | | | | | | | | | | gga Gly | | | 624 |
| _ | _ | | | _ | | | | | | | | | | gct Ala | | | 672 |
| | | | | | | | | • | | | | | | ctg Leu | | | 720 |
| _ | | | | | | | | | | | | | | tgg Trp 255 | | - | 768 |
| _ | | | | | | | | | | | | | | agc Ser | | | 816 |
| _ | Ser | | _ | _ | _ | | | | | | | | | agc Ser | | | 864 |
| | | | | | | | | | | | | | | gag Glu | | | 912 |
| acg | ccc | cca | aat | gac | tgc | cag | caa | gaa | ctg | aat | cta | gtg | agg | act | gtg | | 960 |

| Thr 305 | Pro | Pro | Asn | Asp | Cys 310 | Gln | Gin | Glu | Leu | Asn 315 | Leu | Val | Arg | Thr | Va I 320 | |
|------------|-----|-----|-----|-----|-------------------|-----|-----|-----|-----|------------|-----|-----|-----|-----|-------------|------|
| | | | - | | gtc Val | | | | | | | | | | | 1008 |
| | | | | | ctg Leu | | | | | | | | | | | 1056 |
| | | | - | | ggg Gly | | | | | | | | | | | 1104 |
| | | | | | ttt Phe | | | | | | | | | | | 1152 |
| _ | | | | | cga Arg 390 | | | | | | | | | | | 1200 |
| | | | | | aac Asn | | | | | | | | | | | 1248 |
| | | | | Gln | cag Gin | | | | Leu | | | | | | | 1296 |
| | | | Ala | | cct Pro | | | Pro | | | | | | | | 1344 |
| gtg | cct | gcc | cct | ggg | agc | cag | gtc | tgc | ctc | cag | cct | ctc | agc | tcc | gcc | 1392 |

| Val | Pro 450 | Ala | Pro | Gly | Ser | GIn 455 | Val | Cys | Leu | Gln | Pro 460 | L.eu | Ser | Ser | Ala | |
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| | | | | | agt Ser 470 | | | | | | | | | | | 1440 |
| | | | | | gag Glu | | | | | | | | | _ | | 1488 |
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| | | | | | agc Ser | | | | | - | | _ | | | | 1584 |
| | | | | | gag Glu | | | | | | | | | | | 1632 |
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Ser Ser Gln Cys Leu Leu Glu Tyr Leu Leu Asn Arg Leu Asp Ser Ser 50 55 60

Ser Gly His Val Lys Leu Lys Val Leu Lys Ile Leu Leu Tyr Leu Cys 75 75 80

Gly His Gly Ser Ser Ser Phe Leu Leu IIe Leu Arg Arg Asn Ser Ala 85 90 95

Leu lle Gin Glu Ala Thr Ala Phe Ser Gly Pro Pro Asp Pro Leu His 100 105 110

Gly Asn Ser Leu Tyr Gln Lys Val Arg Ala Ala Ala Gln Asp Leu Gly
115 120 125

Ser Thr Leu Phe Ser Asp Ala Val Pro Gln Pro Pro Ser Gln Pro Pro

130 135 140

Gln lle Pro Pro Pro Ala Gly Met Gly Ala Gln Ala Arg Pro Leu Ser 145 150 155 160

Ala Leu Gin Gly Phe Gly Tyr Thr Lys Glu Ser Ser Arg Thr Gly Ser 165 170 175

Ala Gly Glu Thr Phe Leu Ser Thr IIe Gln Arg Ala Ala Glu Val Val 180 185 190

Ala Asn Ala Val Arg Pro Gly Pro Asp Asn Pro Cys Thr Lys Gly Pro 195 200 205

Leu Pro Tyr Gly Asp Ser Tyr Gln Pro Ala Val Thr Pro Ser Ala Ser 210 215 220

His Thr His Pro Asn Pro Gly Asn Leu Leu Pro Gly Ala IIe Leu Gly 225 230 235 240

Ala Arg Ala Val Arg His Gln Pro Gly Gln Ala Gly Gly Gly Trp Asp 245 250 255

Glu Leu Asp Ser Ser Pro Ser Ser Gln Asn Ser Ser Cys Thr Ser Asn 260 265 270

Leu Ser Arg Ala Ser Asp Ser Gly Ser Arg Ser Gly Ser Asp Ser His

275 280 285

Ser Gly Thr Ser Arg Glu Pro Gly Asp Leu Ala Glu Arg Ala Glu Ala 290 295 300

Thr Pro Pro Asn Asp Cys Gln Gln Glu Leu Asn Leu Val Arg Thr Val 305 310 315 320

Thr Gln Gly Pro Arg Val Phe Leu Ser Arg Glu Glu Thr Gln His Phe 325 330 335

lle Lys Glu Cys Gly Leu Leu Asn Cys Glu Ala Val Leu Glu Leu Leu 340 345 350

Leu Arg Gin Leu Val Gly Thr Ser Glu Cys Glu Gin Met Arg Ala Leu 355 360 365

Cys Ala IIe Ala Ser Phe Gly Ser Ala Asp Leu Leu Pro Gln Glu His 370 375 380

Val Leu Leu Cys Arg Gln Gln Leu Gln Glu Leu Gly Ala Gly Ser 385 390 395 400

Pro Gly Pro Val Thr Asn Lys Ala Thr Lys lle Leu Arg His Phe Glu 405 410 415

Ala Ser Cys Gly Gln Gln Leu Pro Thr Leu Arg Leu Cys Ala Gln Pro

420 425 430

Asn Ser Ala Ala Pro Val Gly Pro Ala Asp Leu Leu Thr Ser Pro 435 440 445

Val Pro Ala Pro Gly Ser Gln Val Cys Leu Gln Pro Leu Ser Ser Ala 450 455 460

Thr Val Val Pro Arg Ser Pro Val Leu Phe Pro Ser Pro Asn Thr Leu 465 470 475 480

Pro Pro Ser Ala Leu Glu Glu Pro Ser Glu Val Arg Thr Gln Leu Val 485 490 495

Cys Ser Ser Glu Gln Gly Thr Glu Ser Glu Gln Arg Leu Glu Asn Thr 500 505 510

Asp Thr Pro Glu Asp Ser Ser Ser Pro Leu Pro Trp Ser Pro Asn Ser 515 520 525

Leu Phe Ala Gly Met Glu Leu Val Ala Cys Pro Arg Leu Pro Cys His 530 535 540

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| | • | | 85 - | | | 90 | | | 95 | | |
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| ttc Phe | | | | | | | | | | | 336 |
| | | _ | | cag Gln | | | | | | | 384 |
| | | | | gac Asp | | | | | | | 432 |
| _ | | | | gcc Ala 150 | | | | | | | 480 |
| _ | | | | ttc Phe | | | | | | | 528 |
| _ | | | Ala | ttc Phe | | | | | | • | 576 |
| | | | | cgc Arg | | | | | | | 624 |
| | | | | gac Asp | | | | | | | 672 |

agc cac ggt ccc cca acc ctg ggg aac cta ctc ccc ggg gcc att cca Ser His Gly Pro Pro Thr Leu Gly Asn Leu Leu Pro Gly Ala lle Pro

| 225 | | | | | 230 | | | | | 235 | | | | | 240 | • |
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| gat Asp | gag Glu | ctg Leu | gac Asp 260 | agc Ser | ggc Gly | ccc Pro | agc Ser | tct Ser 265 | cag Gln | aat Asn | tcc. Ser | tcc Ser | cag Gln 270 | aac Asn | agc Ser | 816 |
| gac Asp | ctg Leu | agc Ser 275 | agg Arg | gtc Val | tcg Ser | gac Asp | tcg Ser 280 | ggc Gly | agt Ser | cat His | tcc Ser | ggc Gly 285 | agc Ser | gac Asp | agc Ser | 864 |
| cat His | tca Ser 290 | ggg Gly | gcc Ala | agc Ser | cgg Arg | gag Glu 295 | ccg Pro | ggt Gly | gac Asp | ctg Leu | gca Ala 300 | gaa Glu | agg Arg | gtc Val | gag Glu | 912 |
| gtg Val 305 | gtg Val | gcc Ala | ctg Leu | agt Ser | gac Asp 310 | tgt Cys | cag Gln | cag Gln | gag Glu | ttg Leu 315 | agc Ser | ttg Leu | gtg Val | agg Arg | act Thr 320 | 960 |
| | act Thr | | | | Arg | | | | | Arg | | | | | | 1008 |
| | atc lle | | | Cys | | | | | Cys | | | | | Gln | | 1056 |
| ct: Lei | g acc ı Thr | tgc Cys 355 | His | ctg Leu | cgt Arg | ggg Gly | acc Thr | Ser | gaa Glu | tgc Cys | acg Thr | cag Gln 365 | Leu | agg Arg | gcg Ala | 1104 |
| | g tgt | | | | | | | | | | | | | | | 1152 |

cac atc ctc cgc acc cgg ccg tgg ctg cag gag ctc agc atg ggc His lie Leu Leu Arg Thr Arg Pro Trp Leu Gin Glu Leu Ser Met Gly ago cog gga cot gtg aco aac aag goo aco aag ato ctg agg cac ttt Ser Pro Gly Pro Val Thr Asn Lys Ala Thr Lys Ile Leu Arg His Phe gag gcc tcc tgt ggg cag ctg tcc cct gcc cgg ggc acc tca gct gag Glu Ala Ser Cys Gly Gln Leu Ser Pro Ala Arg Gly Thr Ser Ala Glu cct ggc ccc aca gcc gcc ctc cca ggc cca tct gac ctg ctg acc gac Pro Gly Pro Thr Ala Ala Leu Pro Gly Pro Ser Asp Leu Leu Thr Asp gct gtg cct ctc cct ggg agc cag gtc ttc ctg cag cct ctg agt tca Ala Val Pro Leu Pro Gly Ser Gln Val Phe Leu Gln Pro Leu Ser Ser acc ccg gtc tcg tcc cgg agc cct gct ccc tca tct ggg atg ccg tcc Thr Pro Val Ser Ser Arg Ser Pro Ala Pro Ser Ser Gly Met Pro Ser ago cot gtg coc aco coa coc coa gat goo too coc att coa goo coc Ser Pro Val Pro Thr Pro Pro Pro Asp Ala Ser Pro Ile Pro Ala Pro

aga cct gaa cgg atc ccg ggg ggc acg gac agc cca aag aga ggc ccc 1584 Arg Pro Glu Arg lle Pro Gly Gly Thr Asp Ser Pro Lys Arg Gly Pro

gga gac ccc agc gag gcc gag gcc aga ctg gca gaa agc agg cgg tgg

Gly Asp Pro Ser Glu Ala Glu Ala Arg Leu Ala Glu Ser Arg Arg Trp

520

525

515 agc agc tgt gcg tgg agc cgc gac tcc ttg ttt gct ggc atg gag ctg 1632 Ser Ser Cys Ala Trp Ser Arg Asp Ser Leu Phe Ala Gly Met Glu Leu 540 535 530 1680 Val Ala Cys Pro Arg Leu Val Gly Ala Gly Ala Ala Ala Gly Glu Ser 555 560 545 550 tgt cct gat gct ccc cgc gcc ccc caa aca tcg tcc cag agg aca gca 1728 Cys Pro Asp Ala Pro Arg Ala Pro Gln Thr Ser Ser Gln Arg Thr Ala 565 570 575 gcc aaa gag cct cct ggc tca gag ccg tca gct ttc gcg ttc ctg aac 1776 Ala Lys Glu Pro Pro Gly Ser Glu Pro Ser Ala Phe Ala Phe Leu Asn 585 590 580 1782 gcc tga Ala <210> <211> 593 <212> PRT <213> Homo sapiens <400> Met Ala Ala Ala Pro Pro Leu Arg Asp Arg Leu Ser Phe Leu His Arg 15 5 . 10 1 Leu Pro IIe Leu Leu Lys Gly Thr Ser Asp Asp Val Pro Cys Pro

25

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Ser Ser Gln Cys Leu Leu Glu Tyr Leu Leu Ser Arg Leu His Ser Ser 50 55 60

Ser Gly His Gly Lys Leu Lys Val Leu Lys Ile Leu Leu Tyr Leu Cys
65 70 75 80

Ser His Gly Ser Ser Phe Phe Leu Leu Ile Leu Lys Arg Asn Ser Ala 85 90 95

Phe IIe Gin Giu Ala Ala Ala Phe Ala Gly Pro Pro Asp Pro Leu His 100 105 110

Gly Asn Ser Leu Tyr Gln Lys Val Arg Ala Ala Ala Gln Asp Leu Gly 115 120 125

Ser Thr Leu Phe Ser Asp Thr Val Leu Pro Leu Ala Pro Ser Gln Pro 130 135 140

Leu Gly Thr Pro Pro Ala Thr Gly Met Gly Ser Gln Ala Arg Pro His
145 150 155 160

Ser Thr Leu Gln Gly Phe Gly Tyr Ser Lys Glu His Gly Arg Thr Gly
165 170 175

Ser Ala Gly Glu Ala Phe Leu Ser Thr lle Gln Lys Ala Ala Glu Val 180 185 190

Val Ala Ser Ala Met Arg Pro Gly Pro Glu Ser Pro Ser Thr Arg Arg 195 200 205

Leu Leu Pro Arg Gly Asp Thr Tyr Gln Pro Ala Met Met Pro Ser Ala 210 215 220

Ser His Gly Pro Pro Thr Leu Gly Asn Leu Leu Pro Gly Ala Ile Pro 225 230 235 240

Gly Pro Arg Ala Val Arg His Gln Pro Gly Gln Ala Gly Gly Gly Trp 245 250 255

Asp Glu Leu Asp Ser Gly Pro Ser Ser Gln Asn Ser Ser Gln Asn Ser 260 265 270

Asp Leu Ser Arg Val Ser Asp Ser Gly Ser His Ser Gly Ser Asp Ser 275 280 285

His Ser Gly Ala Ser Arg Glu Pro Gly Asp Leu Ala Glu Arg Val Glu 290 295 300

Val Val Ala Leu Ser Asp Cys Gln Gln Glu Leu Ser Leu Val Arg Thr 305 310 315 320 Val Thr Arg Gly Pro Arg Ala Phe Leu Ser Arg Glu Glu Ala Gln His 325 330 - 335

Phe lle Lys Ala Cys Gly Leu Leu Asn Cys Glu Ala Val Leu Gln Leu 340 345 350

Leu Thr Cys His Leu Arg Gly Thr Ser Glu Cys Thr Gln Leu Arg Ala 355 360 365

Leu Cys Ala IIe Ala Ser Leu Gly Ser Ser Asp Leu Leu Pro Gln Glu 370 375 380

His lle Leu Leu Arg Thr Arg Pro Trp Leu Gln Glu Leu Ser Met Gly 385 390 395 400

Ser Pro Gly Pro Val Thr Asn Lys Ala Thr Lys Ile Leu Arg His Phe 405 410 415

Glu Ala Ser Cys Gly Gln Leu Ser Pro Ala Arg Gly Thr Ser Ala Glu 420 425 430

Pro Gly Pro Thr Ala Ala Leu Pro Gly Pro Ser Asp Leu Leu Thr Asp 435 440 445

Ala Val Pro Leu Pro Gly Ser Gln Val Phe Leu Gln Pro Leu Ser Ser 450 455 460 Thr Pro Val Ser Ser Arg Ser Pro Ala Pro Ser Ser Gly Met Pro Ser 465 470 475 480

Ser Pro Val Pro Thr Pro Pro Pro Asp Ala Ser Pro Ile Pro Ala Pro 485 490 495

Gly Asp Pro Ser Glu Ala Glu Ala Arg Leu Ala Glu Ser Arg Arg Trp 500 505 510

Arg Pro Glu Arg IIe Pro Gly Gly Thr Asp Ser Pro Lys Arg Gly Pro 515 520 525

Ser Ser Cys Ala Trp Ser Arg Asp Ser Leu Phe Ala Gly Met Glu Leu 530 535 540

Val Ala Cys Pro Arg Leu Val Gly Ala Gly Ala Ala Ala Gly Glu Ser 545 550 555 560

Cys Pro Asp Ala Pro Arg Ala Pro Gln Thr Ser Ser Gln Arg Thr Ala 565 570 575

Ala Lys Glu Pro Pro Gly Ser Glu Pro Ser Ala Phe Ala Phe Leu Asn 580 585 590

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<223>
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| | | |
| С | | 61 |
| | | |
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| <220> | D | |
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| | cttg accaaacctc tggcgaagaa gtccaaagct tcactcgcgg atgctggc | 58 |
| cggugu | oreg adottation tageographic processing to the contract of the | |
| | | |
| <210> | 9 | |
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| cgcgtt | tgga atcactacag gg | 22 |
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| Laatao | guot ouotatuggs | |
| | | |
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| ٠ | <213> | Artificial | |
| | (000) | | |
| | <220> | Description of Autificial Companyation autificially | |
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| | aaagtg | gaga ttgttgccat | 20 |
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| | | | |
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| | (220) | synthesized primer sequence | |
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| | ttgact | gtgc cgttgaatt | 19 |
| | | | |
| | | | |
| | <210> | 18 | |
| | <211> | 17 | |
| | <212> | DNA | |
| | <213> | Artificial | |
| | Z000\ | | |
| | <220> | Description of Autificial Commence on this is in the | |
| | <223> | Description of Artificial Sequence:an artificially | |

24/27

synthesized primer sequence

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|----------------|----------------------------------------------------|----|
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| ggcgtt | cagg aacgcgaaag | 20 |
| | | |
| Z010\ | | |
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| \213/ | ALLITICIAL | |
| <220> | • | |
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| | synthesized primer sequence | |
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| | | |
| <210> | 21 | |
| <211> | 21 | |
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| | Artificial | |
| \L U/ | πιτιτοιαι | |

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| | synthesized primer sequence | |
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| | • | |
| <210> | 22 | |
| <211> | 21 | |
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| | Artificial | |
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| | synthesized primer sequence | |
| <400> | 22 | |
| • | gatg cgatacccgg g | 21 |
| agoreg | Park observable P | |
| | | |
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| <211> | 19 | |
| <212> | DNA | |
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| \223/ | Description of Artificial Sequence:an artificially synthesized primer sequence | |
| | Synthesized primer sequence | |
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| tttgca | gggc tggcaagcc | 19 |
| | | |
| | | |
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| <211> | 39 | |
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| | Artificial | |
| <220> | | |
| <223> | Description of Artificial Sequence an artificially synthesized primer sequence | |
| <400> | 25 | |
| | accet etgegaacga tgaagttett ggeettee | 38 |
| • | | |
| <210> | 26 | |
| <211> | 30 | |
| <212> | DNA : | |
| <213> | Artificial | |
| <220> | | |
| <223> | Description of Artificial Sequence:an artificially synthesized primer sequence | |
| <400> | 26 | |
| gggac | tacaa ggacgatgac gataagtagc | 30 |

<210> 27

<211> 36

<212> DNA

<213> Artificial

<220>

<400> 27

ggccgctact tatcgtcatc gtccttgtag tcccgc